Amendments to the Claims

Please amend the following claims:

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Claim 1. (Previously Amended) A compound of Formula (I):

$$R_2$$
 R_3
 R_3
 R_4
 R_3
 R_4
Formula (I)

wherein

Y is selected from the group consisting of a bond, -C(0), -C(0)O-, -C(0)NE- and $-SO_2-$;

 R_1 is selected from the group consisting of R_7 and R_8 ;

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 R_2 , R_3 , R_4 and R_5 are independently selected from the group consisting of a bond, hydrogen and C_{1-8} alkyl; wherein C_{1-8} alkyl is optionally substituted with one to three substituents independently selected from R_9 , provided that R_2 , R_3 , R_4 or R_5 can only be a bond when forming a monocyclic ring wherein the following monocyclic rings may be formed from R_2 , R_3 , R_4 and R_5 ;

when R_2 and R_3 comprise a bond and C_{1-8} alkyl or optionally when both R_2 and R_3 are C_{1-8} alkyl , R_2 and R_3 together with the atoms to which each is attached will form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R₃ and R₄ comprise a bond and C₁₋₈alkyl or optionally when both R₃ and R₄ are C₁₋₈alkyl, R₃ and R₄ together with the atoms to which each is attached will form a five to seven membered monecyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_3 and R_5 comprise a bond and C_{1-8} alkyl or optionally when both R_3 and R_5 are C_{1-8} alkyl, R_3 and R_5 together with the atoms to which each is attached will form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_4 and R_5 comprise a bond and C_{1-8} alkyl, or optionally when both R_5 and R_5 are C_{1-8} alkyl, R_4 and R_5 together with the atoms to which each is attached will form a four to seven membered monocyclic ring optionally containing one

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to two additional heteroatoms independently selected from the group consisting of N, O and S;

- R_7 , R_8 R_{10} and R_{14} are independently selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkoxy, C₁₋₈alkylcarbonyl, C₁₋₈alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, $N-(C_{1-8}alkyl)$ amino, $N, N-(C_{1-8}dialkyl)$ amino, $-CF_3$ and -OCF3; wherein cycloalkyl/and heterocyclyl are optionally substituted with one to three oxo substituents; and, wherein the aryl and heteroaryl/substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, C1-salkyl, C2-salkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl) amino, $N, N-(C_{1-8} \text{dialkyl}) \text{ ami} \neq 0, -CF_3 \text{ and } -OCF_3;$
- R_8 , R_{12} , R_{13} and R_{17} are independently selected from the group consisting of C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, and $(halo)_{1-3}(C_{1-8})$ alkyl; wherein C_{1-8} alkyl, C_{2-8} alkenyl and C_{2-8} alkynyl are optionally substituted on a terminal carbon

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with one to three substituents independently selected from R_{14} ;

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- R_{11} is selected from the group consisting of hydrogen and C_{1-8} alkyl;
- A is C₁₋₄alkylene optionally substituted with one to two substituents independently selected from R₁₃;
- when R_3 is C_{1-8} alkyl, optionally A and R_3 together with the atoms to which each is attached may form a five to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;
- when R₄ is C₁₋₈alkyl, optionally A and R₄ together with the atoms which each is attached may form a five to seven membered monocyclic ring optionally containing one additional heteroatom selected from the group consisting of N, O and S;
- when R₅ is C₁₋₈alkyl, optionally A and R₅ together with the atoms which each is attached may form a three to seven membered monocyclic ring optionally containing one to two heteroatoms independently selected from the group consisting of N, O and S; and
- B₁ and B₂ are independently selected from the group consisting of C₁₋₂alkylene and C₂alkenylene optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C₁₋₈)alkyl, hydroxy(C₁₋₈)alkoxy, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₁;

and pharmaceutically acceptable salts, racemic mixtures, diastereomers and enantiomers thereof.

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Claim 2. (Original) The compound of claim 1 wherein Y is selected from the group consisting of -C(0) and $-SO_2$.

Claim 3. (Original) The compound of claim 1 wherein Y is selected from $-SO_2-$.

Claim 4. (Original) The compound of claim 1 wherein R_1 is selected from R_7 .

Claim 5. (Original) The compound of claim 1 wherein R_2 , R_3 , R_4 and R_5 are independently selected from the group consisting of hydrogen and C_{1-4} alkyl.

Claim 6. (Original) The compound of claim 1 wherein R_2 , R_3 , R_4 and R_5 are independently selected from the group consisting of hydrogen and methyl.

Claim 7. (Original) The compound of claim 1 wherein R_6 is optionally present and is one to three substituents independently selected from the group consisting of halogen, C_{1-8} alkoxy, R_{10} , R_{12} , $-N(R_{11})C(O)-R_{10}$, $-N(R_{11})C(O)-R_{12}$, $-N(R_{11})SO_2-R_{10}$, $-N(R_{11})C(O)-N(R_{11},R_{12})$, $-N(R_{11})C(O)-N(R_{12},R_{17})$, $-OC(O)-N(R_{11},R_{12})$, $-OC(O)-N(R_{12},R_{17})$, $-OC(O)-R_{10}$ and $R_{10}-(C_{1-8})$ alkoxy.

Claim 8. (Original) The compound of claim 1 wherein R_{ϵ} is optionally present and is one to three substituents independently selected from the group consisting of halogen,

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Claim 9. (Original) The compound of claim 1 wherein R_6 is optionally present and is one to two substituents independently selected from the group consisting of R_{10} , $-N(R_{11})C(0)-R_{10}$, $-N(R_{11})C(0)-N(R_{11},R_{12})$, $-N(R_{11})C(0)-N(R_{12},R_{17})$, $-OC(0)-N(R_{11},R_{12})$, $-OC(0)-N(R_{12},R_{17})$ and R_{10} -methoxy.

Claim 10. (Original) The compound of claim 1 wherein R₇ is selected from the group consisting of aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, C₁₋₈alkylcarbonyl, C₁₋₈alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, N-(C₁₋₈alkyl)amino, N, N-(C₁₋₆dialkyl)amino, -EF₃ and -OCF₃; and, wherein the aryl and heteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N, N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃.

Claim 11. (Original) The compound of claim 1 wherein R₁₀ is selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₁₋₈alkoxy, C₁₋₈alkoxycarbonyl, carboxyl, arylcarbonyl, arylsulfonyl, -CF₃ and -OCF₃; wherein cycloalkyl and heterocyclyl are optionally substituted with one to three oxe substituents; and, wherein the aryl portion

of the arylcarbonyl substituent is optionally substituted with one to five substituents independently selected from C_{1-8} alkoxy.

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Claim 12. (Original) The compound of (claim 1 wherein R₁₀ is selected from the group consisting of cyclopropyl, 1,3dihydro-2H-isoindolyl, 2-azabicyclo[2/.2.2]octyl, piperidinyl, morpholinyl, phenyl, naphthalenyl, thienyl, 1H-pyrrolyl and pyridinyl; wherein cyclopropyl, pigeridinyl, morpholinyl, phenyl, naphthalenyl, thienyl, 1H-pyrrolyl and pyridinyl are optionally substituted with one to four substituents independently selected from the group consisting of chlorine, fluorine, bromine, methyl, isopropyl, t-butyl, methoxy, tbutoxycarbonyl, carboxyl, phenylcarbonyl, -CF3 and -OCF3; wherein 1,3-dihydro-2H-isoindolyl is optionally substituted with oxo; wherein 2-azabicycl/p[2.2.2]octyl is optionally substituted with phenylsulfo yl, and, wherein the phenyl portion of the phenylcarbon 1 substituent is optionally substituted with one to two substituents independently selected from methoxy.

Claim 13. (Original) The compound of claim 1 wherein R_{12} is selected from the group consisting of C_{1-8} alkyl and C_{2-8} alkynyl optionally substituted on a terminal carbon with R_{14} .

Claim 14. (Original) The compound of claim 1 wherein R_{12} is selected from the group consisting of C_{1-4} alkyl and C_{2-4} alkynyl optionally substituted on a terminal carbon with R_{14} .

Claim 15. (Original) The compound of claim 1 wherein R_{12} is selected from the group consisting of t-butyl and ethynyl; wherein ethynyl is optionally substituted on a terminal carbon with a substituent independently selected from R_{14} .

Claim 16. (Original) The compound of claim I wherein R_{14} is selected from the group consisting of aryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkoxy, C_{1-8} alkylcarbonyl, C_{1-8} alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino $N-(C_{1-8}$ alkyl)amino, N_1 - N_2 - N_3 - N_4 - N_3 - N_4 - N_3 - N_4 -N

Claim 17. (Original) The compound of claim 1 wherein R_{11} is selected from the group consisting of hydrogen and C_{1-4} alkyl.

Claim 18. (Original) The compound of claim 1 wherein R_{11} is hydrogen.

Claim 19. (Original) The compound of claim 1 wherein A is selected from the group consisting of methylene and ethylene.

Claim 20. Canceled

Claim 21. (Original) The compound of claim 1 wherein B_1 and B_2 are independently selected from the group consisting of $-CH_2-$, $-(CH_3)_3-$ and $-(CH)_2-$ optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C_{1-4}) alkyl, hydroxy(C_{1-4}) alkexy, C_{1-4} alkyl, C_{2-4} alkeyl, C_{2-4} alkynyl,

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 C_{1-4} alkoxy, carboxyl, amino, $N-(C_{1-4}$ alkyl) amino, $N, N-(C_{1-4}$ dialkyl) amino, $-CF_3$ and $-OCF_3$.

Claim 22. (Original) The compound of claim 1 wherein B_1 is selected from the group consisting of $-CH_2-$, $-(CH_2)_2-$ and $-(CH)_2-$ optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C_{1-4}) alkyl, hydroxy(C_{1-4}) alkoxy, C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{1-4} alkoxy, carboxyl, amino, $N-(C_{1-4}$ alkyl) amino, $N,N-(C_{1-4}$ dialkyl) amino, $-CF_3$ and $-OCF_3$; and, wherein, B_2 is selected from $-(CH_2)_2-$.

Claim 23. (Original) The compound of claim 1 wherein B_1 is selected from the group consisting of $-CH_2-$, $-(CH_2)_2-$ and $-(CH)_2-$.

Claim 24. (Original) The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:

$$R_{6}$$
 R_{6}
 R_{7}
 R_{8}
 R_{1}
 R_{2}
 R_{3}
 R_{3}

wherein B_1 , R_1 , R_3 , R_5 , A and R_6 are dependently selected from the group consisting of:

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B ₁	R ₁	R ₃	R _S	A	R ₆
(CH ₃) ₂	4- Tol	Н	Н	CH ₂	4-NHC(0)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	4- Tol	н	Н	CH ₂	4-NHC(0) - (2,4,6-Cl ₃) Ph;
(CH ₂) ₃	4- Tol	Н	н	CH3	4-NHC(O)-[2,6-(OMe) ₂]Ph;
CH3	Ph	Н	Н	CH2	4; NHC(O)-(2,6-F ₂)Ph;
(CH ₂) ₃	Ph	н	H	CH₂	4-NHC(O) - (2,6-Cl ₂) Ph;
(CH ₂) ₂	Ph	н	H	CH₂	4-[2,6-(OMe) ₂]Ph;
(CH ₂) ₃	4- Tol	H	Н	CH2	4-NHC(O)-(2-Me)Ph;
(CH ₂) ₃	4- Tol	Н	H	CH ₂	4-NHC(0)-(2-C1)Ph;
(CH ₂) ₂	4- Tol	н	н	CH ₂	4-NHC(0)-(2,6-F ₂)Ph;
(CH ₂) ₂	4- Tol	Ħ	н .	CH ₂	4-NHC(O)-(2-CF ₃)Ph;
(CH ₂) ₃	4- Tol	H	н	CH ₂	4-NHC(0)-(2-OCF ₃)Ph;
(CH ₂) ₂	4- Tol	H	H	CH ₂	4-NHC(0)-(2-Br)Ph;
$(CH_2)_2$	Ph	н	Н	CH ₂	4-NHC(0)-(2,6-F ₂)Ph;
CH ₂	Ph	н //	H	CH ₂	4-NHC(0)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	4- Tol	Н	н	CH2	4-[2,6-(OMe) ₂]Ph;
CH ₂	Ph	H	н	CH ₂	4-NHC(0)-[2,6-(OMe) ₂]·Ph;
(CH ₂) ₂	4- Tol	Н	Н	CH ₂	4-CC-(4-t-butyl)Ph;
(CH ₃) ₂	4 -	Н	. н	CH ₂	4-CC-Ph;
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End

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	Tol				
(CH ₃) ₂	4- Tol	н	Н	CH ₂	4-NHC(O)-Ph;
(CH ₂) ₂	4- Tol	Н	H	CH ₂	4-NHC(O) - [4-C(O) - [2,5- (OMe) ₂] Ph] Ph;
(CH ₂) ₂	4- Tol	Н	Н	CH ₂	4-NHC(O)-CH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) 3	Ph	H	Н	CH ₂	4-NHC(O)-NH-(2,6-Cl ₂)Ph;
(CH ₂) ₃	Ph	H	н	CH ₃	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	4- Tol	H	H	CH ₂	4-OCH ₃ -Ph;
$(CH_2)_2$	4 -	Н	Н	CH₂	4-NHC(0) - (2,4,6-
	Tol				isopropyl3) Ph;
(CH ₂) ₂	4 <i>-</i> Tol	Н	Н	CH ₂	4-(1 <i>H</i> -pyrrol-1-yl);
(CH ₂) ₂	4- Tol	Н	н	ĊH₂	4-Ph;
(CH ₂) ₂	Ph	н	н	CH ₂	4-NHC(0)-NH-(2,6-F ₂)Ph;
(CH ₂) ₂	4- Tol	Н	н	CH ₂	3-NHC(O)-(2,6-F ₂)Ph;
(CH ₂) ₂	4- Tol	Н	н/	CH ₂	3-NHC(O) - [2,6-(OMe) ₂] Ph;
(CH ₃) ₃	4- Tol	Н	н	CH2	3-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	н	CH ₃	CH ₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₃	Ph	CĦ³	Н	CH ₂	4-NHC(0)-(2,6-Cl ₂)Ph;
(CH) ₂	Ph	H	Н	CH3	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	Ή	Н	CH ₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH) ₂	Ph	н	н	CH ₂	4-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph /	H	Н	CH ₂	4-(2,4,6-F ₃)Ph;

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(CH ₂) ₂	Ph	H	Н	CH ₂	4-(2,3,5,6-F ₄)Ph;
(CH ₂) ₂	Ph	н	Н	CH2	4-0-t-butoxy;
$(CH_2)_2$	Ph	H	H	(CH ₂) ₂	;
(CH ₂) ₃	Ph	н	Н	CH ₂	4-(#,3-dihydro-1,3-dioxo-
					2Hfisoindol-2-yl);
(CH ²) ²	Ph	н	H	\mathtt{CH}_2	4/-NHC(O) - (2-CO ₂ H) Ph;
(CH ₂) ₂	Ph	н	H	CH ₂	4-(2,5-diMe-1H-pyrrol-1-
				//	/yl);
(CH ₂) ₂	Ph	Н	н	CH ₂	4-NHC(0)-4-pyridinyl;
(CH ₃) ₂	Ph	н	H	CH ₂	4-NHSO ₂ -(2,6-Cl ₃)Ph;
(CH ₂) ₂	Ph	н	н	CH,	4-OC(O)-N(CH ₃) ₂ ;
(CH ₂) ₂	Ph	Н	Н	CH2	4-NHC(O)-(1-t-
•	•			//	butoxycarbonyl)4-
		•	/	//	piperidinyl;
(CH ₃) ₃	4 -	н	н //	CH2	4-NHC(0)-(2,6-Cl ₂)Ph;
	FPh				
(CH ₂) ₂	4 -	Н	# #	CH ₂	4-NHC(O)-[2,6-(OMe) ₂]Ph;
	FPh				
(CH ₂) ₂	Ph	н	н	CH ₂	4-OC(O)-4-morpholinyl;
$(CH_2)_2$	Ph	н	н	CH ₃	4-OC(O)N(1so-propyl)2;
(CH ₂) ₂	Ph	н//	Н	CH ₂	4-t-butyl;
(CH ₂) ₂	Ph	H	Н	CH₂	4-NHC(0)-4-piperidinyl;
(CH ₂) ₂	Ph /	н	H	CH ₂	4-NHC(0)-(3,5-Cl ₂)4-
					pyridinyl;
(CH ₂) ₂	Ph	Н	н	CH ₂	4-NHC(O)-NMe ₂ ;
(CH ₂) ₂	Ph	н	H	CH ₂	3-F-4-[OCH ₂ (2,6-Cl ₃)Ph];
(CH ₂) ₂	2/-	H	Н	CH ₂	4-OC(O)-NMe2;
	Thi				
(CH ₂) ₂	Ph	н	н	CH ₂	4-NHC(0)-t-butyl;
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(CH ₂) ₂	Ph	н	H	CH ₂	4-NHC(O)-(2-OMe)1-
					naphthalenyl;
$(CH_2)_2$	2-	н	Н	CH_2	4-NHC(0)-(2,6-Cl2)Ph;
	Thi				
$(CH_2)_2$	Ph	Н	Н	CH ₂	4-NHC(0)-cyclopropyl;
$(CH_2)_2$	Ph	Н	Н	CH ₃	4-NHC(0)-(2,2,3,3-
					Me4)cyclopropyl;
(CH ₃) ₂	Ph	Н	Н	CH₂	A-NHC(O)-iso-propyl;
(CH ₂) ₂	Ph	H	Н	CH_2	4-NHC(0)-(2-SO ₂ Ph)-2-
					azabicyclo[2.2.2]oct-3-yl;
(CH ₂) ₂	2-	Н	H	CH ₂	4-NHC(O)-(3,5-Cl ₂)4-
	Thi				pyridinyl;
(CH ₃) ₂	Ph	Н	Н	CH₂	4-NHC(0)-(2-
					Me) cyclopropyl;
$(CH_2)_2$	Ph	Н	H	ДСН₃	4-(2,6-diMe)Ph;
$(CH_2)_2$	Ph	Н	н	CH3	4-(2,6-Cl ₂)Ph;
(CH ₂) ₂	2-	Н	н	CH2	4-(2,6-Cl ₂) Ph;
	Thi				
(CH ₂) ₂	2-	H	/H	CH_2	4-(2,6-diMe)Ph;
	Thi				
(CH ₂) ₃	2 -	н	/ н	CH₃	4-[2,6-(OMe) ₂]Ph;
	Thi				
(CH ₂) ₂	2-	H	H	CH_2	4-(4-fluoro-1,3-dihydro-
	Thi				1,3-dioxo-2H-isoindol-2-
	,				yl);
(CH ₂) ₂	2-	H	H	CH ₂	4-NHC(O)-NMe ₂ ;
	Thi				
(CH ₂) ₂	2 -	H	H	CH ₂	$4-OC(O)-NMe_2;$
	Thi				
(CH ₂) ₂	24	Н	H	CH ₂	4-OC(0)-(4-morpholinyl);
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	Thi				/
(CH ₂) ₂	2-	н	H	CH ₂	4-0C(0)-(4-Me-1-
	Thi				piperazinyl);
$(CH_2)_2$	Ph	Н	н	CH2	4-OC(O)-(4-Me-1-
			٠		piperazinyl);
$(CH_2)_3$	Ph	H	Ħ	CH ₂	4-N (Me) C(O) - (2,6-Cl ₂) Ph;
(CH ₂) 2	Ph	H	н	CH ₂	4-N(Me)C(O)-(3,5-Cl ₂)4-
					pyridinyl;
(CH ₃) ₂	2 -	Н	н	CH ₂	4-N(Me)C(O)-(3,5-Cl ₃)4-
	Thi				pyridinyl;
$(CH_2)_2$	2-	н	Н	CH ³	4-N(Me)C(O)-(2,6-Cl ₂)Ph;
	Thi				
(CH ₃) ₃	2 -	Н	Н	CH₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
	Thi			I	
(CH ₂) ₃	2 -	H	н //	CH2	4-(1,3-dihydro-1,3-dioxo-
	Thi		J -		2H-isoindol-2-yl);
(CH3)3	Ph	н	H	CH3	4-(1,3-dihydro-4,7-
		l:			dimethyl-1,3-dioxo-2H-
					isoindol-2-yl);
$(CH_2)_2$	2-	н	Н	CH ₂	4-(1,3-dihydro-4,7-
	Thi				dimethyl-1,3-dioxo-2H-
	. ,				isoindol-2-yl);
CH ₂	2- //	H	Н	CH ₂	4-NHC(O)-(3,5-Cl ₂)4-
	Thi				pyridinyl;
CH ₂	2-//	H	H	CH ₂	4-NHC(O)-(2,6-Cl ₂)Ph;
	Thi				
(CH ₂) ₂	Ph	H	H	CH ₂	4-(1,1-dioxido-3-oxo-1,2-
	7				<pre>benzisothiazol-2(3H)-yl);</pre>
(CH ₃) ₃	Ph	н	н	CH ₂	4-(4-chloro-1,3-dihydro-
/					1,3-dioxo-2H-isoindol-2-



y1);

and,

 $(CH_2)_2$ Ph CH_2

4-(7.9-dioxo-8-

azaspiro[4.5]dec-8-yl);

and pharmaceutically acceptable salts, racemic mixtures, diastereomers and enantiomers thereof.

Claim 25. (Previously Amended) A compound having Formula (II):

 R_6 Formula (II)

wherein

Y is selected f_x^{M} om the group consisting of -C(0) - and -SO₂-;

 R_1 is selected from the group consisting of R_7 and R_8 ;

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R₃, R₃, R₄ and R₅ are independently selected from the group consisting of a bond, hydrogen and Cl₋₈alkyl; wherein C₁₋₈alkyl is optionally substituted with one to three substituents independently selected from R₉; provided that R₂, R₃, R₄ and R₅ can only be a bond when forming a monocylic ring wherein the following monocylic rings may be formed from R₃, R₃, R₄ and R₅:

when R₂ and R₃ comprise a bond and C₁₋₈alkyl or optionally when both R₂ and R₃ are C₁₋₈alkyl, R₂ and R₃ together with the atoms to which each are attached form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_3 and R_4 comprise a bond and C_{1-8} alkyl or optionally when both R_3 and R_4 are C_{1-8} alkyl, R_3 and R_4 together with the atoms to which each are attached form a five to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_3 and R_5 comprise a bond and C_{1-8} alkyl or optionally when both R_3 and R_5 are C_{1-8} alkyl, R_3 and R_5 together with the atoms to which each are attached form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_4 and R_5 comprise a bond and C_{1-8} alkyl or optionally when both R_4 and R_5 are C_{1-8} alkyl, R_4 and R_5 together with the atoms to which each are attached form a four to seven membered monocyclic ring optionally containing one to two

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additional heteroatoms independently selected from the group consisting of N, O and S;

- R_7 R_9 , R_{10} and R_{14} are independently selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkoxy, C_{1-g} alkylcarbonyl, C_{1-g} alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, $N-(C_{1-8}alkyl)$ amino, $N-(C_{1-8}dialkyl)$ amino, $-CF_3$ and -OCF3; wherein cycloalkyl and heterocyclyl are optionally substituted with one to three oxo substituents; and, wherein the aryl and meteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, C1-salkyl, C_{2-8} alkenyl, C_{2-8} alk $\frac{1}{2}$ nyl, C_{1-8} alkoxy, carboxyl, amino, $N-(C_{1-8}alkyl)$ amino $\iint N, N-(C_{1-8}dialkyl)$ amino, $-CF_3$ and $-OCF_3$;
- R_8 , R_{12} , R_{13} and R_{17} are independently selected from the group consisting of C_1 alkyl, C_2 alkenyl, C_2 alkynyl, and $(halo)_{1-3}(C_{1-8})$ alkyl; wherein C_{1-8} alkyl, C_2 alkenyl and C_2 alkynyl are optionally substituted on a terminal carbon

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with one to three substituents independently selected from R_{14} ;

 R_{11} is selected from the group consisting of hydrogen and C_{1-8} alkyl;

A is C_{1.4}alkylene optionally substituted with one to two substituents independently selected from R₁₃;

when R₃ is C₁₋₈alkyl, optionally A and R₃ together with the atoms to which each is attached form a five to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_4 is C_{1-8} alkyl, optionally A and R_4 together with the atoms to which each is attached form a five to seven membered monocyclic ring optionally containing one additional heteroatom selected from the group consisting of N, O and S;

when R₅ is C₁₋₈alkyl, optionally A and R₃ together with the atoms to which each is attached form a three to seven membered monocyclic ring optionally containing one to two heteroatoms independently selected from the group consisting of N, O and S;

B is selected from the group consisting of C₁₋₂alkylene and C₂alkenylene optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C₁₋₈)alkyl, hydroxy(C₁₋₈)alkoxy, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃; and,

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n is an integer from 1 to 2;

and pharmaceutically acceptable salts racemic mixtures, diastereomers and enantiomers thereof.

Claim 26. (Previously Amended) A process for preparing a compound of Formula (III):

wherein

 R_1 is selected from the group consisting of R_7 and R_8 ;

 R_{1} , R_{10} , and R_{14} are independently selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl

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optionally substituted with one to five substituents independently selected from the group consisting of halogen, C_{1-8} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-8} alkoxy. C_{1-8} alkylcarbonyl, C_{1-8} alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, $N-(C_{1-8}$ alkyl)amino, $N,N-(C_{1-8}$ dialkyl)amino, $-CF_3$ and $-OCF_3$; wherein cycloalkyl and heterocyclyl are optionally substituted with one to three oxe substituents; and, wherein the aryl and heteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently substituted with one to five substituents independently selected from the group consisting of halogen, C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkoxy, carboxyl, amino, $N-(C_{1-8}$ alkyl)amino, $N,N-(C_{1-8}$ dialkyl)amino, $-CF_3$ and $-OCF_3$;

 R_8 , R_{13} and R_{17} are independently selected from the group consisting of C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, and $(halo)_{1-3}(C_{1-8})$ alkyl; wherein C_{1-8} alkyl, C_{2-8} alkenyl and C_{2-8} alkynyl are optionally substituted on a terminal carbon with one to three substituents independently selected from R_{14} ;

 R_{150} is selected from the group consisting of hydroxy, amino, NO_2 and $R_6\,;$

R₆ is optionally present and is one to three substituents independently selected from the group consisting of halogen, $C_{1\text{-8}}alkoxy, \ R_{10}, \ R_{12}, -N(R_{11})C(O) - R_{10}, -N(R_{11})C(O) - R_{12}, \\ -N(R_{11})SO_3 - R_{10}, -N(R_{11})SO_2 - R_{12}, -N(R_{11})C(O) - N(R_{11}, R_{10}), \\ -N(R_{11})C(O) - N(R_{11}, R_{12}), -N(R_{11})C(O) - N(R_{12}, R_{17}), -C(O) - N(R_{11}, R_{10}), \\ -C(O) - N(R_{12}, R_{17}), -C(O) - N(R_{11}, R_{12}), -OC(O) - N(R_{11}, R_{10}), \\ -OC(O) - N(R_{11}, R_{12}), -OC(O) - N(R_{12}, R_{17}), -OC(O) - R_{10}, -OC(O) - R_{12}, \\ -O-R_{10} \ and \ R_{10}-(C_{1-8}) \ alkoxy;$

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 R_{11} is selected from the group consisting of hydrogen and C_{1-8} alkyl; and,

B₁ and B₂ are independently selected from the group consisting of C₁₋₂alkylene and C₂alkenylene optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C₁₋₈)alkyl, hydroxy(C₁₋₈)alkoxy, C₁₋₈alkyl, C₁₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃;

and pharmaceutically acceptable salts, racemic mixtures, diastereomers and enantiomers thereof;

comprising reacting a compound of Formula (IV)

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wherein

R₁₆ is selected from the group consisting of halogen, mixed anhydride and hydroxy;

with a compound of Formula (V)

R₁₅
OMe
O • HCl
Formula (V);

in the presence of appropriate coupling agents, bases and solvents to form the compound of Formula (II).

Claim 27. (Original) The process of claim 25 wherein R_{15} is selected from the group consisting of hydroxy, iodine, bromine and NO_2

Claim 28. (Original) The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:

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ON OH OH

Claim 29. (Original) The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:

Claim 30. (Original) The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:

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CH₃

Claim 31. (Original) The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:

Claim 32. (Original) The compound of claim 1 wherein the compounds are effective antagonists of an integrin receptor.

Claim 33. (Original) The compound of claim 32 wherein the compound is a selective antagonist of an $\alpha 4$ integrin receptor.

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Claim 34. (Original) The compound of claim 33 wherein the $\alpha4$ integrin receptor is selected from the group consisting of the $\alpha4\beta1$ and $\alpha4\beta7$ integrin receptor.

Claim 35. (Original) The compound of claim 32 wherein the compound is an antagonist of at least two $\alpha 4$ integrin receptors.

Claim 36. (Original) The compound of claim 35 wherein the two $\alpha4$ integrin receptors are selected from the group consisting of the $\alpha4\beta1$ and $\alpha4\beta7$ integrin receptor.

Claims 37-43 (Canceled)

Claim 44. (Original) A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable carrier.

Claim 45. (Original) A pharmaceutical composition made by mixing a compound of claim 1 and a pharmaceutically acceptable carrier.

Claim 46. (Original) A method for the treatment of an integrin mediated disorder ameliorated by inhibition of an $\alpha 4$ integrin receptor comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

Claim 47. (Canceled)

Claim 48. (Original) The method of claim 47 wherein the $\alpha 4$ integrin receptor is selected from the group consisting of the $\alpha 4\beta 1$ and $\alpha 4\beta 7$ integrin receptor.

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Claim 49. (Original) The method of claim 46 wherein the compound inhibiting the $\alpha4$ integrin receptor is selected from the group consisting of a selective antagonist of the $\alpha4\beta1$ integrin receptor, a selective antagonist of the $\alpha4\beta7$ integrin receptor and an antagonist of the $\alpha4\beta1$ and $\alpha4\beta7$ integrin receptors.

Claim 50. (Original) The method of claim 46 wherein the integrin mediated disorder is selected from the group consisting of inflammatory disorders, autoimmune disorders and cell-proliferative disorders.

Claim 51. (Original) The method of claim 46 wherein the integrin mediated disorder is selected from the group consisting of inflammation disorders, autoimmunity disorders, asthma, bronchoconstriction, restenosis, atherosclerosis, psoriasis, rheumatoid arthritis, inflammatory bowel disease, irritable bowel disease, irritable bowel disease irritable bowel syndrome, transplant rejection and multiple sclerosis.

Claim 52. (Currently Amended) The <u>method compound</u> of claim 46 wherein the integrin mediated disorder is selected from the group consisting of asthma, bronchoconstriction, restenosis, atherosclerosis, psoriasis, rheumatoid arthritis, inflammatory bowel disease, irritable bowel disease, irritable bowel syndrome, transplant rejection and multiple sclerosis.

Claim 53. (Currently Amended) The <u>method compound</u> of claim 46 wherein the integrin mediated disorder is selected from the group consisting of asthma, bronchoconstriction, restenosis, atherosclerosis, irritable bowel syndrome and multiple sclerosis.

Claim 54. (Original) The method of claim 46 wherein the therapeutically effective amount of the compound of claim 1 is from about 0.01 mg/kg/day to about 300 mg/kg/day.

Claim 55. (Currently Amended) The method of claim 46 further comprising administering to a subject in need thereof a therapeutically effective amount of the pharmaceutical composition of claim 44 compound of claim 1 and a pharmaceutically acceptable excipient.

Claim 56. (Currently Amended) The method of claim 55 wherein the therapeutically effective amount of the pharmaceutical composition of claim 44 compound of claim 1 and a pharmaceutically acceptable excipient is from about 0.01 mg/kg/day to about 300 mg/kg/day.

Claim 57. (Origina) The compound of claim 1 wherein R_7 is selected from the group consisting tolyl, phenyl and thienyl.

Claim 58. (Currently Amended) The method of claim 46 wherein the integrin mediated disorder is a cell-proliferation disorders disorders.

